DOCKET NO.: ISIS-5425 PATENT

Application No.: Not Yet Assigned

Preliminary Amendment - First Action Not Yet Received

This listing of claims will replace all prior versions, and listings, of claims in the application.

## **Listing of Claims:**

Please cancel claims 1-44 and insert new claims 45-50 as presented below.

45 (New). A compound comprising a plurality of linked nucleosides, wherein: at least one of the nucleosides is a 2'-deoxy'-2'-fluoro, 2'-O-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-O-C<sub>2</sub>-C<sub>20</sub>-alkenyl, 2'-O-C<sub>2</sub>-C<sub>20</sub>-alkynyl, 2'-S-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-S-C<sub>2</sub>-C<sub>20</sub>-alkenyl, 2'-NH-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-NH-C<sub>2</sub>-C<sub>20</sub>-alkynyl nucleoside; and

at least one of the nucleosides is a 5' terminal nucleoside having a non-aromatic lipophilic molecule linked to the 5'-position of the nucleoside.

46 (New). The compound of claim 45 wherein at least one of said nucleosides is a 2'-deoxy-2'-fluoro or a 2'-methoxy nucleoside.

47 (New). A compound comprising a plurality of linked nucleosides, wherein: at least one of the nucleosides is a 2'-deoxy'-2'-fluoro, 2'-O-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-O-C<sub>2</sub>-C<sub>20</sub>-alkenyl, 2'-O-C<sub>2</sub>-C<sub>20</sub>-alkynyl, 2'-S-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-S-C<sub>2</sub>-C<sub>20</sub>-alkenyl, 2'-NH-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-NH-C<sub>2</sub>-C<sub>20</sub>-alkynyl nucleoside: and

at least one of the nucleosides is a 5' terminal nucleoside having a non-aromatic molecule linked to the 5'-position of the nucleoside.

48 (New). The compound of claim 47 wherein at least one of said nucleosides is a 2'-deoxy-2'-fluoro or a 2'-methoxy nucleoside.

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49 (New). A compound comprising a plurality of linked nucleosides, wherein: at least one of the nucleosides is a 2'-deoxy'-2'-fluoro, 2'-O-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-O-C<sub>2</sub>-C<sub>20</sub>-alkynyl, 2'-S-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-S-C<sub>2</sub>-C<sub>20</sub>-alkenyl, 2'-NH-C<sub>1</sub>-C<sub>20</sub>-alkyl, 2'-NH-C<sub>2</sub>-C<sub>20</sub>-alkynyl nucleoside; and

at least one of the nucleosides is a 5' terminal nucleoside having a lipophilic molecule linked to the 5'-position of the nucleoside.

50 (New). The compound of claim 49 wherein at least one of said nucleosides is a 2'-deoxy-2'-fluoro or a 2'-methoxy nucleoside.